Applicant: Richard F Selden et al.

Serial No.: 09/686,497 Filed: October 11, 2000

Page : 2 of 7

Amendments to the Claims:

This listing of claims replaces all prior versions and listings of claims in the application:

•NO.6050---P. 3•

Aπomey's Docket No.: 10278-022001 / 0020 (98-6 CIP)

Listing of Claims:

1. (Currently Amended) A synthetic nucleic acid sequence that encodes <u>human</u> α-galactosidase, wherein at least one non-common codon or less-common codon has been replaced by a common codon and wherein the synthetic nucleic acid has one or more of the following properties:

it has a continuous stretch of at least 150 codons all of which are common codons; it has a continuous stretch of common codons, which continuous stretch comprises at least 33% 60% of the codons of the synthetic nucleic acid sequence;

at least 94% of the codons in the sequence encoding the protein are common codons, wherein by a common codon is meant Ala (gcc); Arg (cgc); Asn (aac); Asp (gac); Cys (tgc); Gln (cag); Gly (ggc); His (cac); Ile (atc); Leu (ctg); Lys (aag); Pro (ccc); Phe (ttc); Ser (agc); Thr (acc); Tyr (tac); Glu (gag); Val (gtg), Met (atg) and Trp (tgg).

- 2. (Original) The synthetic nucleic acid sequence of claim 1, where the α-galactosidase nucleic acid is inserted into a non-transformed cell.
- 3. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the number of non-common or less-common codons remaining is less than 15.
- 4. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the number of non-common or less-common codons remaining, taken together, are equal or less than 6% of the codons in the synthetic nucleic acid sequence.
- 5. (Original) The synthetic nucleic acid sequence of claim 1, wherein all non-common or less-common codons are replaced with common codons.

Applicant: Richard F Selden et al.

Serial No.: 09/686,497 Filed: October 11, 2000

Page : 3 of 7

Attorney's Docket No.: 10278-022001 / 0020 (98-6 CIP)

- 6. (Original) The synthetic nucleic acid sequence of claim 1, wherein at least 96% of the codons in the synthetic nucleic acid sequence are common codons.
- 7. (Original) The synthetic nucleic acid sequence of claim 1, wherein at least 98% of the codons in the synthetic nucleic acid sequence are common codons.
- 8. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein all the codons of the synthetic nucleic acid sequence are common codons.
 - 9. (Original) A vector comprising the synthetic nucleic acid sequence of claim 1.
 - 10. (Original) A cell comprising the nucleic acid sequence of claim 1.
- 11. (Original) A method of producing α-galactosidase comprising culturing the cell of claim 10 under conditions in which the nucleic acid is expressed.
- 12. (Currently Amended) A method for preparing a synthetic nucleic acid sequence encoding <u>human</u> α-galactosidase, comprising:

identifying a non-common codon and a less-common codon in a non-optimized gene sequence which encodes an α -galactosidase protein; and

replacing at least 94% of the non-common and less-common codons with a common codon encoding the same amino acid as the replaced codon,

wherein by a common codon is meant: Ala (gcc); Arg (cgc); Asn (aac); Asp (gac); Cys (tgc); Gln (cag); Gly (ggc); His (cac); Ile (atc); Leu (ctg); Lys (aag); Pro (ccc); Phe (ttc); Ser (agc); Thr (acc); Tyr (tac); Glu (gag); Val (gtg), Met (atg) and Trp (tgg).

13. (Original) The method of claim 12, wherein at least 96% of the non-common and less-common codons are replaced with a common codon encoding the same amino acid as the replaced codon.

Attorney's Docker No.: 10278-022001 / 0020 (98-6 CIP)

Applicant: Richard F Selden et al.

Serial No.: 09/686,497

Filed: October 11, 2000

Page : 4 of 7

14. (Original) The method of claim 12, wherein at least 98% of the non-common and less-common codons are replaced with a common codon encoding the same amino acid as the replaced codon.

15-25. (Canceled)

- 26. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the nucleic acid has a continuous stretch of at least 200 common codons.
- 27. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the nucleic acid has a continuous stretch of at least 250 common codons.
- 28. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the nucleic acid has a continuous stretch of at least 300 common codons.
- 29. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the nucleic acid has a continuous stretch of common codons, which continuous stretch comprises at least 70% of the codons of the synthetic nucleic acid sequence.
- 30. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the nucleic acid has a continuous stretch of common codons, which continuous stretch comprises at least 80% of the codons of the synthetic nucleic acid sequence.
- 31. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the nucleic acid has a continuous stretch of common codons, which continuous stretch comprises at least 90% of the codons of the synthetic nucleic acid sequence.
- 32. (Previously Presented) The synthetic nucleic acid sequence of claim 1, wherein the nucleic acid has a continuous stretch of common codons, which continuous stretch comprises at least 95 % of the codons of the synthetic nucleic acid sequence.